

FIGURE 1

## Scheme 1: synthesis of difluorocombretastatin

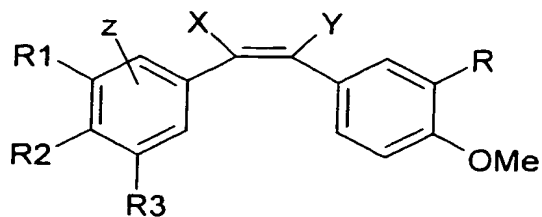
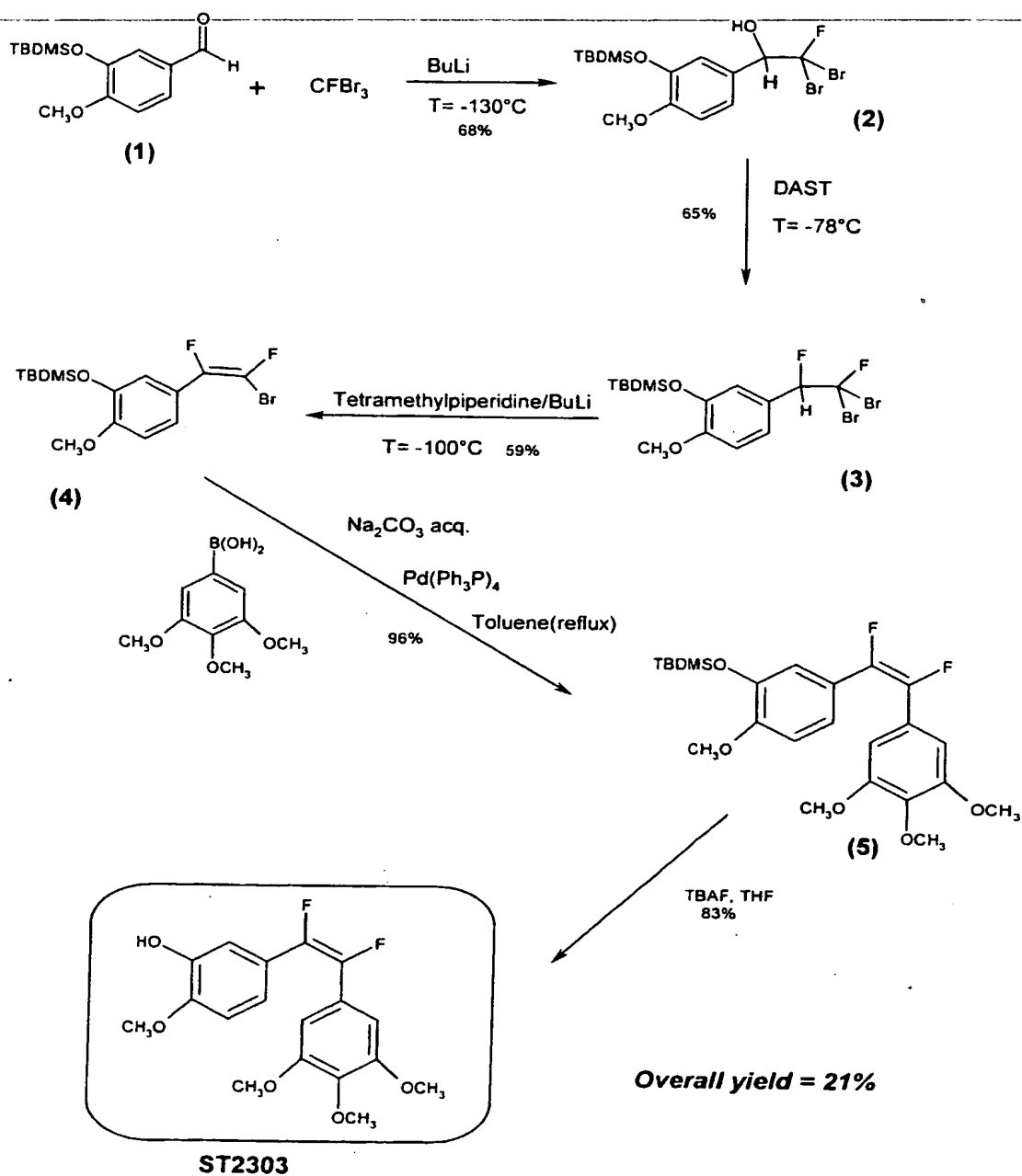


FIGURE 2

Scheme 2: Synthesis of difluoro-Nitro- and difluoro-Amino-combretastatin

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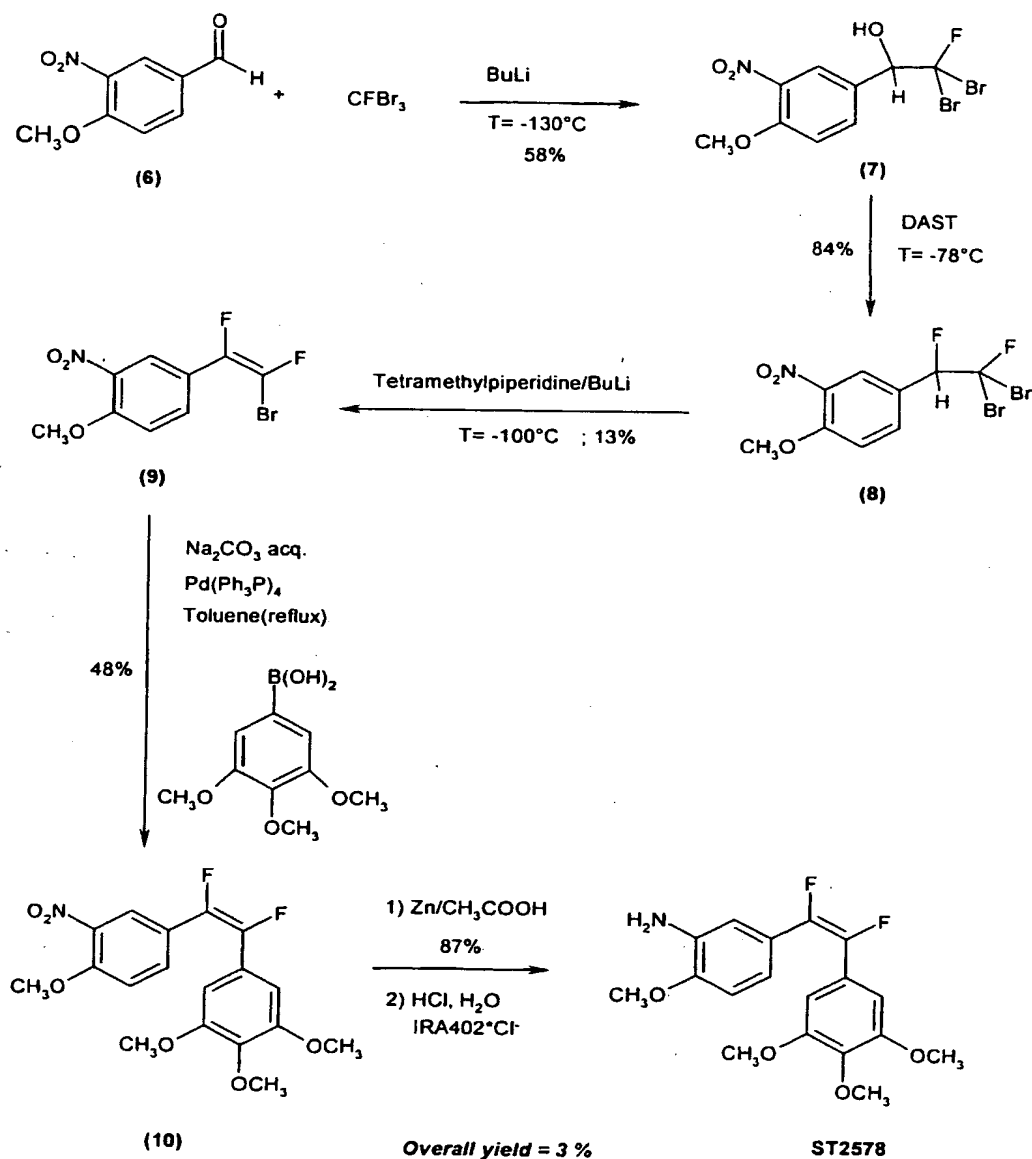
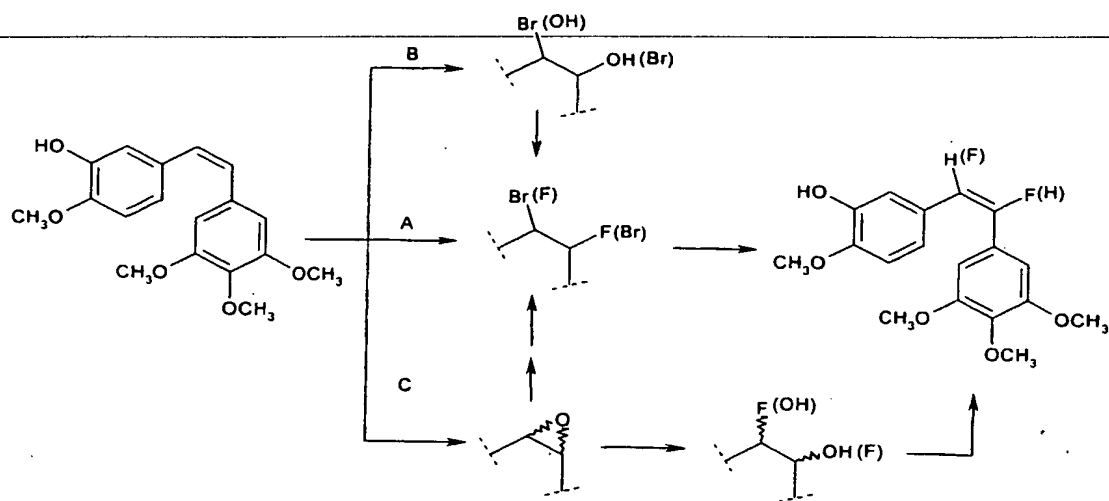


FIGURE 3

## Scheme 3: Synthesis of monofluorocombretastatins



## Scheme 3a: Total synthesis approach to monofluorocombretastatin

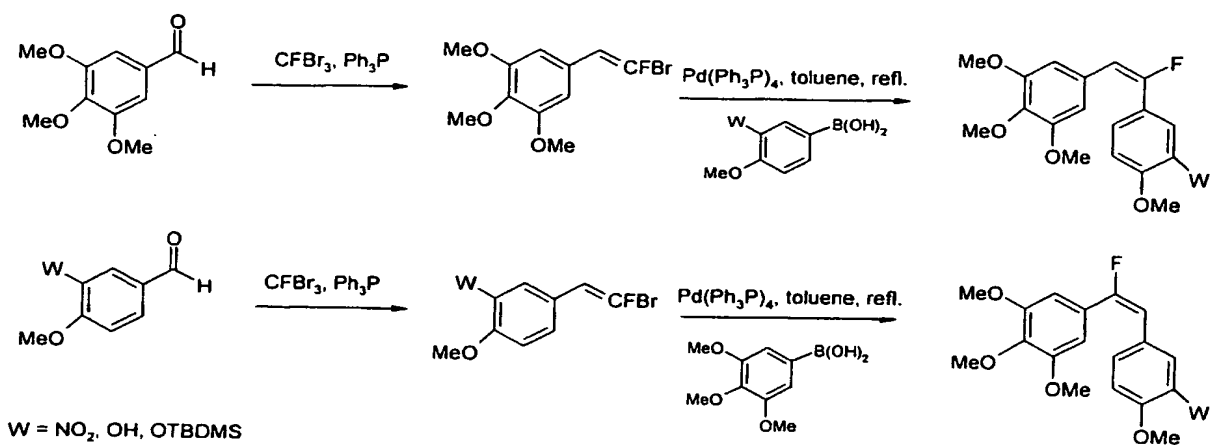


FIGURE 4

Scheme 4: Synthesis of disodium-phosphate prodrug difluoro-combretastatin (ST2493)

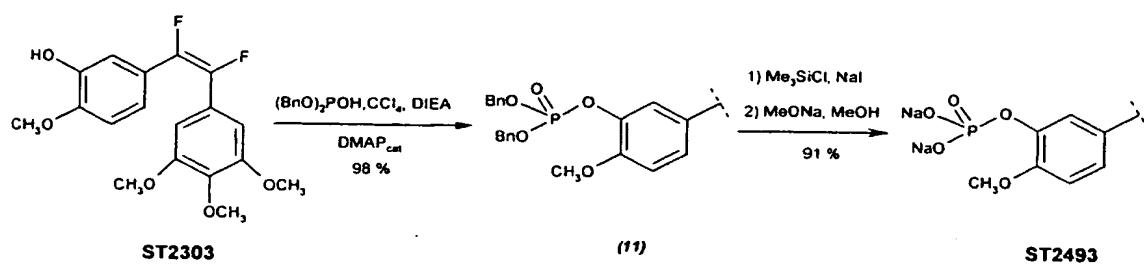


FIGURE 5

Scheme 5: Synthesis of disodium mono-difluorocombretastatin-4-O-methoxyphosphate [12]

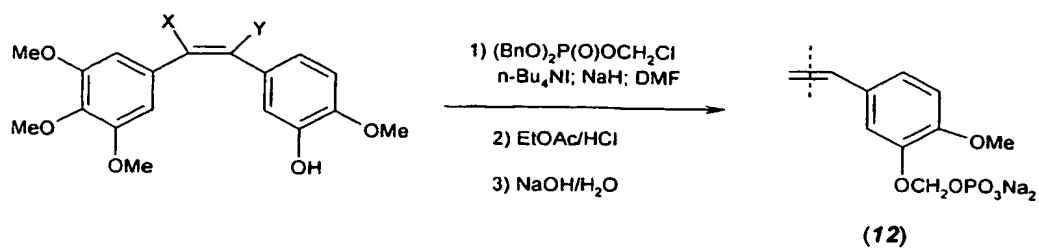


FIGURE 6

Scheme 6: General Procedure for obtaining amide derivatives [13].

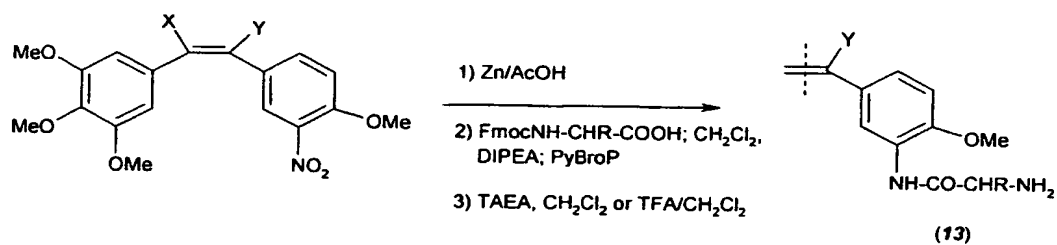


FIGURE 7

## Scheme 7: Synthesis of bromofluorocombretastatin

